THE LANCET Planetary Health

Supplementary appendix

This appendix formed part of the original submission and has been peer reviewed. We post it as supplied by the authors.

Supplement to: Vanker A, Barnett W, Workman L, et al. Early-life exposure to indoor air pollution or tobacco smoke and lower respiratory tract illness and wheezing in African infants: a longitudinal birth cohort study. *Lancet Planet Health* 2017; **1:** e328–36.

Supplemental information

Early-Life Exposure to Indoor Air Pollution or Tobacco Smoke and Lower Respiratory Illness in African Infants.

Aneesa Vanker¹, Whitney Barnett¹, Lesley Workman¹, Polite M. Nduru¹, Peter D. Sly², Robert P. Gie³, Heather J. Zar¹

¹ Department of Paediatrics and Child Health, Red Cross War Memorial Children's Hospital, and MRC Unit on Child & Adolescent Health, University of Cape Town, Klipfontein Road, Rondebosch, 7700, South Africa

² Children's Health and Environment Program, Child Health Research Centre, The University of Queensland 62 Graham St South Brisbane, Queensland, Australia, 4101

³Department of Paediatrics and Child Health, Tygerberg Children's Hospital, Stellenbosch University, Francie van Zijl Avenue, Tygerberg, 7505, South Africa

Supplemental Table 1: Methods

| Study population and proce | edures |
|---|---|
| 1.1 Health Facilities | Mbekweni and Newman health facilities provide free primary health to women and children, including antenatal care, a strong prevention of mother to child transmission (PMTCT) HIV program, childhood immunizations including thirteen valent pneumococcal conjugate vaccine (PCV13) and care for intercurrent illness. Hospital referral is to the single public hospital serving the area, Paarl hospital. |
| 1.2 Sociodemographic | Sociodemographic data were collected using a questionnaire adapted from the South African Stress and Health Study (SASH).(1) A composite SES score was developed based on current employment status and standardised scores of educational level, household income and a composite asset index made up of access to household resources, amenities and market access categorising participants as being lowest SES, low-moderate SES, moderate-high SES or high SES. |
| Measuring exposure to IAP | |
| 1.3 Dwelling categorization | Dwellings were catergorized as a poor structure if there were 2 or less of 6 dwelling dimensions (type of home, building material, water supply, type of toilet, kitchen type and ventilation).(2) An implementation of the Alkire-Foster method, a flexible technique used to incorporate a number of dimensions of poverty or well-being, that can complement poverty assessment (2, 3) was applied to the dwelling characteristics. Six dwelling factors were used; type of home (formal versus informal), primary building material (brick or cement versus other materials), water supply (piped into dwelling or yard), toilet facilities (non communal flush), kitchen type (separate room in house) and ventilation in the kitchen area (pipe or duct to exterior). Dwellings were then categorised according to the number of dimensions lacking. This method defines a dwelling as a "poor structure" if it lacks one-third or more of the factors considered.(4) |
| 1.4 Pollutant measurement | Particulate matter (PM ₁₀) (personal air sampling pump – SKC Aircheck 52 ^R) |
| equipment | Carbon monoxide (CO) (Altair ^R Carbon Monoxide single gas detection unit) |
| | Nitrogen dioxide (NO) and sulphur dioxide (Radiello ^R adsorbent filters in polyethylene diffusive body) |
| | • Volatile organic compounds (VOC) benzene and toluene (Markes ^R thermal desorption tubes using passive diffusion tubes).(4) |
| | All measurements were done in the communal/main living room, away from windows and doors, approximately 1.5 meters from the ground. |
| 1.5 National Ambient Air Quality Standards | Expected exposure for each pollutant based on an averaging period of 1 year for each measure; PM10: 40ug/m³, NO2: 40ug/m³, benzene: 5ug/m³, toluene: 240ug/m³, CO: >30mg/m³ (not more than 88 hours). (5) An average concentration based on the 2-week duration in the home was obtained for sulphur dioxide/nitrogen dioxide and volatile organic compounds; 24-hour averages were obtained for particulate matter. Carbon monoxide data was downloaded to a computer and the frequency of exceedance above the hourly ambient standard was calculated. Based on the 10 minute readings, total hourly concentrations were computed using the trapezium rule. Two consecutive ten minute CO readings were used to represent parallel sides of a trapezium and the 10 minute interval to represent the distance between the parallel sides (width). The trapezium formula; half the sum of the parallel sides multiplied by the width, was then applied. The sum of six consecutive trapezia areas to represent total CO concentration in an hour was then calculated. Using this approach hourly concentrations were then determined for the entire duration of the CO device in the household. (4) |
| Measuring exposure to ETS | |
| 1.6 Self-reported exposure | Maternal tobacco smoking and exposure were assessed using detailed self report questionnaires at enrolment. Post-natal follow-up questionnaires on child respiratory health included questions on tobacco smoke exposure from partners and household members. Maternal smoking was quantified as pack years, where one pack year was defined as 20 cigarettes smoked daily for one year. Maternal nicotine dependence was assessed using the Fagerström test for nicotine dependence, a well-validated questionnaire which scores tobacco dependence as low, low to moderate, moderate or high. The Alcohol, Smoking and Substance Involvement Screening Test (ASSIST) was administered to assess substance use and substance-related risk.(6) |
| 1.7 Urine cotinine | Urine cotinine tests were performed using the IMMULITE ^R 1000 Nicotine Metabolite Kit (Siemens Medical Solutions Diagnostics ^R , Glyn Rhonwy, United Kingdom).(7) |
| measurement | |
| Assessing lower respiratory | |
| 1.8 WHO pneumonia/LRTI case definitions | WHO pneumonia/LRTI case definition: cough or difficulty breathing and age-specific tachypnea or lower chest wall in-drawing).(8) WHO severe pneumonia/LRTI case definition: any child under 2 months of age with signs of pneumonia/LRTI or in a child of any age with danger signs (cyanosed, unable to drink, seizures, or decreased level of consciousness).(8, 9) |
| 1.9 Surveillance for pneumonia/LRTI | Active surveillance for pneumonia/LRTI in the cohort was undertaken as described, using community field workers, a short message system (SMS) phone system, ongoing monitoring of cases at health facilities and study staff who could be contacted by a mother at all times.(10) |

| 1.10 Recognition of LRTI/wheezing | Child caregiver reports at each study visit and episodes identified through the active surveillance for respiratory symptoms associated with LRTI was used to measure wheeze. Study nurses at the primary clinics performed active surveillance and assessed presenting infants in real time. (10, 11) Training of nursing staff included video-clips demonstrating clinical signs. The study doctor provided regular on-site refresher training and competency assessment.(10) |
|-----------------------------------|---|
| Statistical analysis | |
| 1.11 Confounding variables | Potential confounding variables included birth weight, gender, ethnicity (site), SES status, weight for age Z (WAZ) score, (12) maternal HIV status, crowding, household |
| | characteristics, fossil fuel usage, vaccination status, nutritional status and feeding in the first 6 months status. |

Supplemental Table 2: Indoor air pollution (IAP) measurements recorded at antenatal and postnatal visits

| | | Antenat | al | | Postnatal | | | | |
|---|-----------------------------|---------------------------|--------------------------|---------|-----------------------------|---------------------------|--------------------------|---------|--|
| IAP Measure Ambient Concentration | Mbekweni Median (IQR) | Newman Median (IQR) | All Median (IQR) | P Value | Mbekweni Median (IQR) | Newman Median (IQR) | All Median (IQR) | P Value | |
| Particulate Matter (PM10) (ug/m³) | 31·77 (12·36 - 62·73) | 36.04 (13.01 - 65·84) | 33·41 (12·49 - 64·80) | 0.348 | 30·29 (14·67 – 51·05) | 28·44 (10·47 - 53·71) | 29·47 (12·59 - 52·48) | 0.328 | |
| Nitrogen Dioxide (ug/m³) | 6·87 (2·50 - 14·56) | 7·12 (3·84 - 11·28) | 7·03 (3·31 - 12·66) | 0.622 | 6·34 (2·81 - 14·57) | 5·28 (2·48 - 11·25) | 5·83 (2·58 - 12·55) | 0.130 | |
| Sulphur Dioxide (ug/m³) | 0·00 (0·00 - 0·34) | 0·00 (0·00 - 0·17) | 0·00 (0·00 - 0·28) | 0.039 | 0.00 | 0·00 (0·00 - 0·00) | 0·00 (0·00 - 0·00) | 0.794 | |
| Benzene (ug/m³) | 4·50 (1·46 - 17·71) | 3·88 (1·83 - 8·56) | 4·28 (1·74 - 11·39) | 0.637 | 2·81 (0·75 - 14·41) | 3·22 (1·46 - 7·57) | 3·08 (1·06 - 9·46) | 0.312 | |
| Toluene(ug/m³) | 16·06 (5·84 - 42·92) | 17·54 (8·24 - 46·48) | 16·88 (7·04 - 44·57) | 0.213 | 14·72 (4·79 - 48·77) | 15·89 (6·52 - 51·66) | 15·50 (5·90 - 48·97) | 0.286 | |
| Average Carbon Monoxide Per Hour (mg/m³) | 0·00 (0·00 - 3·22) | 0·00 (0·00 - 9·09) | 0·00 (0·00 - 6·21) | 0.105 | 0·00 (0·00 - 0·00) | 0·00 (0·00 - 5·57) | 0·00 (0·00 - 0·00) | 0.015 | |

IAP, indoor air pollution; IQR, inter-quartile range

Supplemental Table 3: Correlation of maternal antenatal cotinine and self-reported total smoke exposure

| | Mbekweni | | Newman | | All | | |
|----------------------------|---------------|---------------|---------------|---------------|---------------|---------------|--|
| | Sensitivity | Specificity | Sensitivity | Specificity | Sensitivity | Specificity | |
| Estimate, % | 71.30 | 59.00 | 95.40 | 35.70 | 83.60 | 53.80 | |
| 95% Confidence Interval, % | 66.80 - 75.50 | 51.70 - 66.00 | 93.00 – 97.10 | 23.40 - 49.60 | 81.00 - 86.00 | 47.40 - 60.10 | |

Supplemental Table 4: Risk factors for lower respiratory tract illness (LRTI) requiring hospitalization

| | | | Antenatal R | isk Factors | | | | | Postnatal R | isk Factors | | |
|---|----------------------|---------------------|----------------------|---------------------|-----------------------|---------|--------------------|---------------------|--------------------|--------------------|-------------------------|---------|
| | | oisodes 524 | | in Model : 245 | | | All Ep N = | | | in Model : 127 | | |
| | Amb., N = 387 | Hsp., N =137 | Amb., N = 177 | Hsp., N = 68 | OR (95%CI) | P value | Amb., N = 387 | Hsp., N =137 | Amb., N = 100 | Hsp, N = 27 | OR (95%CI) | P value |
| Site | | | | | | | | | | | | |
| Mbekweni | 251 (65%) | 73 (53%) | 82 (46%) | 27 (40%) | 0·77 (0·35 - 1·72) | 0.524 | 251 (65%) | 73 (53%) | 50 (50%) | 12 (44%) | 0·40 (0·10 - 1·66) | 0206 |
| Smoke exposure status | s | | | | | | | | | | | |
| Antenatal maternal non- smoker | 75 (19%) | 22 (16%) | 36 (20%) | 14 (20%) | 1 | | | | | | | |
| Antenatal / Postnatal maternal active smoker | 134 (35%) | 61 (45%) | 70 (40%) | 31 (46%) | 0·67 (0·28 - 1·60) | 0.362 | 109 (28%) | 50 (37%) | 33 (33%) | 12 (44%) | 1·12 (0·23 - 5·45) | 0.893 |
| Antenatal maternal passive smoker / Postnatal household smoker | 158 (41%) | 44 (32%) | 71 (40%) | 23 (34%) | 0·58 (0·23 - 1·50) | 0.264 | 269 (70%) | 101 (74%) | 77 (77%) | 21 (78%) | 0·50 (0·12 - 2·19) | 0.359 |
| Unknown | 20 (5%) | 10 (7%) | | | | | | | | | | |
| Infant characteristics | | | | | | | | | | | | |
| Gender (male) | 240 (62%) | 91 (66%) | 123 (69%) | 50 (74%) | 1·05 (0·52 - 2·11) | 0.887 | 240 (62%) | 91 (66%) | 64 (64%) | 20 (74%) | 2·10 (0·64 - 6·97) | 0.223 |
| WAZ at birth | -0·7 (-1·4, -0·1) | -0·8 (-1·5, 0·0) | -0·8 (-1·5, -0·1) | -0·9 (-1·6, 0·0) | 0·64 (0·51 - 0·82) | <0.001 | 0·2 (-0·6, 1·1) | -0·4 (-1·7, 0·8) | 0·6 (-0·3, 1·3) | 0.0 (-0.6, 0.9) | 1·06 (0·60 - 1·89) | 0.836 |
| Maternal HIV exposure | 107 (28%) | 39 (28%) | 22 (12%) | 13 (19%) | 2·04 (0·76 - 5·45) | 0.156 | 107 (28%) | 39 (28%) | 9 (9%) | 7 (26%) | 11·14 (1·71 - 72·73) | 0.012 |
| Age (mid-interval in days) | 4·7 (3 - 7) | 2·5 (1·5 - 7) | 5·2 (3·3 - 7·1) | 2·9 (1·5 - 8·1) | 0·90 (0·82 - 1·00) | 0.050 | 4·7 (3 - 7) | 2·5 (1·5 - 7) | 5·2 (3·3 - 7·1) | 2·3 (1·5 - 8·4) | 0·89 (0·74 - 1·07) | 0.215 |
| SES quartiles (compar | red to high SES) | | | | | | | | | | | |
| Lowest SES | 96 (25%) | 37 (27%) | 47 (27%) | 15 (22%) | 0·56 (0·22 - 1·42) | 0.221 | 96 (25%) | 37 (27%) | 35 (35%) | 10 (37%) | 0·75 (0·21 - 2·65) | 0.659 |
| Low-mod SES | 139 (36%) | 40 (29%) | 64 (36%) | 23 (34%) | 0·99 (0·41 - 2·37) | 0.977 | 61 (16%) | 19 (14%) | 17 (17%) | 5 (19%) | 0·60 (0·14 - 2·53) | 0.490 |
| Mod-high SES | 83 (21%) | 31 (23%) | 36 (20%) | 12 (18%) | 0·75 (0·28 - 1·97) | 0.558 | 74 (19%) | 20 (14%) | 25 (25%) | 3 (11%) | 0·27 (0·05 - 1·34) | 0.109 |
| High SES | 69 (18%) | 29 (21%) | 30 (17%) | 18 (26%) | 1 | | 59 (15%) | 24 (18%) | 23 (23%) | 9 (33%) | | |
| Unknown | | | | | | | 97 (25%) | 37 (27%) | | | | |

| Method of feeding | | | | | | | | | | | | |
|---|------------------------------|--------------------|--------------------|--------------------|------------------------|-------|--------------------|--------------------|--------------------|--------------------|-----------------------|-------|
| Duration of exclusive breast feeding (months) | 2.0 (1.0 - 3·4) | 2.0 (1.0 - 4.0) | 2.0 (1.0 - 3·2) | 1·7 (1.0 - 4.0) | 0.95 (0·79 - 1·14) | 0.577 | 2.0 (1.0 - 3·4) | 2.0 (1.0 - 4.0) | 1·9 (1.0 - 5.0) | 1·5 (1.0 - 3.0) | 0·86 (0·64 - 1·15) | 0.304 |
| Toluene indoor air poll | Toluene indoor air pollution | | | | | | | | | | | |
| Below ambient standard | 223 (58%) | 76 (55%) | 172 (97%) | 61 (90%) | 1 | | 125 (32%) | 41 (30%) | 88 (88%) | 22 (81%) | | |
| Above ambient standard | 8 (2%) | 7 (5%) | 5 (3%) | 7 (10%) | 5·13 (1·43 - 18·36) | 0.012 | 17 (4%) | 5 (4%) | 12 (12%) | 5 (19%) | 1·63 (0·40 - 6·70) | 0.500 |
| Unknown | 156 (40%) | 54 (40%) | | | | | 245 (64%) | 91 (66%) | | | | |

WAZ, weight-for-age z-score; HIV, human immunodeficiency virus; SES, socio-economic status; Amb., ambulatory; Hsp., hospitalized

Supplemental Table 5: Risk factors for lower respiratory tract illness (LRTI) requiring oxygen

| | | | Antenatal R | isk Factors | | | | | Postnatal Ris | sk Factors | | |
|--------------------------------------|----------------------|----------------------|----------------------|---------------------|-----------------------|---------|----------------------|------------------------|----------------------|---------------------|----------------------------|---------|
| | All Ep N = | | N = | in Model 244 | | | N = | pisodes = 521 | | l in Model = 127 | | |
| | No Oxygen N = 452 | Oxygen N = 69 | No Oxygen N = 209 | Oxygen N = 35 | OR (95%CI) | P value | No Oxygen N = 452 | Oxygen N = 69 | No Oxygen N = 116 | Oxygen N = 11 | OR (95%CI) | P value |
| Site | | | | | | • | | | | | | |
| Mbekweni | 285 (63%) | 36 (52%) | 90 (43%) | 18 (51%) | 1·57 (0·49 - 5.00) | 0.446 | 285 (63%) | 36 (52%) | 58 (50%) | 4 (36%) | 0·06 (0·0 - 21·10) | 0.346 |
| Smoke exposure status | | | | | | | | | | | | |
| Antenatal maternal non- smoker | 85 (19%) | 10 (14%) | 43 (21%) | 7 (20%) | 1.0 | | | | | | | |
| Antenatal passive smoker (cotinine) | 178 (39%) | 23 (33%) | 78 (37%) | 15 (43%) | 1·34 (0·36 - 4·92) | 0.659 | | | | | | |
| Antenatal active smoker (cotinine) | 166 (37%) | 29 (42%) | 88 (42%) | 13 (37%) | 1·03 (0·25 - 4·34) | 0.966 | | | | | | |
| Postnatal maternal smoker | | | | | | | 136 (30%) | 23 (33%) | 41 (35%) | 4 (36%) | 1·78 (0·03 - 96·25) | 0.778 |
| Postnatal household smoker | | | | | | | 319 (71%) | 48 (70%) | 90 (78%) | 8 (73%) | 0·28 (0·00 - 31·15) | 0.597 |
| Unknown | 23 (5%) | 7 (10%) | | | | | | | | | | |
| Infant characteristics | | • | | | | | | • | | • | • | |
| Gender (male) | 284 (63%) | 44 (64%) | 149 (71%) | 23 (66%) | 0·72 (0·26 - 1.98) | 0.521 | 284 (63%) | 44 (64%) | 75 (65%) | 9 (82%) | 2·66 (0·06 - 119·69) | 0.614 |
| WAZ at birth | -0·8 (-1·4, -0·1) | -0·5 (-1·4, -0·1) | -0·8 (-1·5, -0·1) | -0·8 (-1·6, 0·1) | 0.96 (0.62 - 1.51) | 0.875 | 0·2 (-0·7, 1·1) | -0·25 (-1·77, 0·73) | 0·6 (-0·5, 1·3) | 0·2 (0·0, 0·9) | 1·12 (0·22 - 5·85) | 0.892 |
| Maternal HIV exposure | 127 (28%) | 17 (25%) | 28 (13%) | 6 (17%) | 1·54 (0·35 - 6·79) | 0.568 | 127 (28%) | 17 (25%) | 13 (11%) | 3 (27%) | 308·86 (0·02 - 5853189) | 0.254 |
| Age (mid-interval in days) | 4·7 (2·9 - 7·1) | 2·3 (1·1 - 4·2) | 5·2 (2·9 - 7·4) | 2·5 (1·5 - 3·4) | 0·76 (0·62 - 0·93) | 0.006 | 4·7 (2·9 - 7·1) | 2·35 (1·13 - 4·19) | 5·2 (3·3 - 7·5) | 2·0 (0·7 - 2·8) | 0·48 (0·18 - 1·28) | 0.142 |
| SES quartiles (compare | ed to high SES) | | | | | _ | | _ | | | | |
| Lowest SES | 121 (27%) | 12 (17%) | 58 (28%) | 4 (11%) | 0·33 (0·07 - 1·59) | 0.168 | 117 (26%) | 16 (23%) | 43 (37%) | 2 (18%) | 1·20 (0·01, 100·91) | 0.935 |
| Low-mod SES | 154 (34%) | 24 (35%) | 74 (35%) | 12 (34%) | 1·00 (0·27 - 3·71) | 0.994 | 65 (14%) | 15 (22%) | 18 (16%) | 4 (36%) | 6·16 (0·05, 699·62) | 0.451 |
| Mod-high SES | 94 (21%) | 19 (28%) | 38 (18%) | 10 (29%) | 1·35 (0·33 - 5·58) | 0.677 | 81 (18%) | 12 (17%) | 25 (22%) | 3 (27%) | 2·45 (0·09, 155·99) | 0.672 |
| High SES | 83 (18%) | 14 (20%) | 39 (19%) | 9 (26%) | 1 | | 74 (16%) | 9 (13%) | 30 (26%) | 2 (18%) | , | |

| Unknown | | | | | | | 115 (25%) | 17 (25%) | | | | |
|---|--------------------|--------------------|-------------------|-------------------|-------------------------|-------|----------------|----------------------|-------------------|-------------------|----------------------|-------|
| Method of feeding | | | | | | | | | | | | |
| Duration of exclusive breast feeding (months) | 2·0 (1·0 - 3·5) | 2·1 (1·0 - 3·6) | 1·8 (1·0 -3·2) | 2.0 $(1.0 - 4.0)$ | 1·10 (0·84 - 1·45) | 0.470 | 4.0 (2.4, 9.0) | 3·64 (3·00, 9·14) | 4·0 (2·0, 9·1) | 3·0 (3·0, 8·9) | 0·51 (0·13, 2·01) | 0.337 |
| Toluene indoor air pol | lution | | | | | | | | | | | |
| Below ambient standard | 263 (58%) | 34 (49%) | 203 (97%) | 29 (83%) | 1 | | | | | | | |
| Above ambient standard | 9 (2%) | 6 (9%) | 6 (3%) | 6 (17%) | 13·21 (1·96 - 89·16) | 0.008 | 22 (5%) | 0 (0%) | 17 (15%) | 0 (0%) | 1·00 (0·98, 1·01) | 0.514 |
| Unknown | 180 (40%) | 29 (42%) | | | | | | | · | | | |

WAZ, weight-for-age z-score; HIV, human immunodeficiency virus; SES, socio-economic status

Supplemental Table 6: Multivariate analysis for lower respiratory tract infection (LRTI) and postnatal environmental exposures

| | IRR (95% CI) | P value |
|--|---|---|
| Site | Hut (55% C1) | 1 value |
| Mbekweni | 1.23 (0.89 - 1.70) | 0.207 |
| Smoke exposure | 123 (0 0) 1 (0) | 0 20, |
| Maternal self-report smoking | 1.22 (0.89 - 1.70) | 0.221 |
| Any household member self-report smoking | 0.94 (0.68 - 1.30) | 0.712 |
| Infant characteristics | 0 71 (0 00 1 20) | 0 / 12 |
| Male | 1.67 (1.30 - 2.15) | 0.000 |
| WAZ at birth | 0.93 (0.83 - 1.04) | 0.226 |
| Maternal HIV exposure | 1.31 (0.90 - 1.91) | 0.160 |
| Age in months | 0.90 (0.88 - 0.93) | 0.000 |
| SES quartiles | | * *** |
| Lowest SES | 1.26 (0.87 - 1.83) | 0.218 |
| Low-mod SES | 1.63 (1.15 - 2.33) | 0.006 |
| Mod-high SES | 0.98 (0.68 - 1.43) | 0.925 |
| High SES | 1 | |
| Method of feeding | | |
| Duration of exclusive breast feeding (months) | | |
| Duration of exclusive breast feeding (months) | 0.97 (0.91 - 1.04) | 0.361 |
| Indoor Air Pollutant Exposure | 0.97 (0.91 - 1.04) | 0.361 |
| | | |
| Indoor Air Pollutant Exposure (N = 429) | 0·97 (0·91 - 1·04) IRR (95% CI) | 0·361 P value |
| Indoor Air Pollutant Exposure (N = 429) Site | IRR (95% CI) | P value |
| Indoor Air Pollutant Exposure (N = 429) Site Mbekweni | | |
| Indoor Air Pollutant Exposure (N = 429) Site Mbekweni Indoor air pollutant exposure | IRR (95% CI) 0.91 (0.61 - 1.37) | P value 0-661 |
| Indoor Air Pollutant Exposure (N = 429) Site Mbekweni Indoor air pollutant exposure PM10 above ambient standard | IRR (95% CI) | P value |
| Indoor Air Pollutant Exposure (N = 429) Site Mbekweni Indoor air pollutant exposure | IRR (95% CI) 0.91 (0.61 - 1.37) 0.61 (0.33 - 1.12) | P value 0.661 0.110 |
| Indoor Air Pollutant Exposure (N = 429) Site Mbekweni Indoor air pollutant exposure PM10 above ambient standard Infant characteristics | IRR (95% CI) 0.91 (0.61 - 1.37) 0.61 (0.33 - 1.12) 1.73 (1.18 - 2.52) | P value 0.661 0.110 0.005 |
| Indoor Air Pollutant Exposure (N = 429) Site Mbekweni Indoor air pollutant exposure PM10 above ambient standard Infant characteristics Male WAZ at birth | IRR (95% CI) 0.91 (0.61 - 1.37) 0.61 (0.33 - 1.12) 1.73 (1.18 - 2.52) 0.94 (0.79 - 1.13) | P value 0·661 0·110 0·005 0·523 |
| Indoor Air Pollutant Exposure (N = 429) Site Mbekweni Indoor air pollutant exposure PM10 above ambient standard Infant characteristics Male WAZ at birth Maternal HIV exposure | IRR (95% CI) 0.91 (0.61 - 1.37) 0.61 (0.33 - 1.12) 1.73 (1.18 - 2.52) 0.94 (0.79 - 1.13) 1.43 (0.76 - 2.70) | P value 0·661 0·110 0·005 0·523 0·226 |
| Indoor Air Pollutant Exposure (N = 429) Site Mbekweni Indoor air pollutant exposure PM10 above ambient standard Infant characteristics Male WAZ at birth Maternal HIV exposure Age in months | IRR (95% CI) 0.91 (0.61 - 1.37) 0.61 (0.33 - 1.12) 1.73 (1.18 - 2.52) 0.94 (0.79 - 1.13) | P value 0·661 0·110 0·005 0·523 |
| Indoor Air Pollutant Exposure (N = 429) Site Mbekweni Indoor air pollutant exposure PM10 above ambient standard Infant characteristics Male WAZ at birth Maternal HIV exposure Age in months SES Quartiles | IRR (95% CI) 0.91 (0.61 - 1.37) 0.61 (0.33 - 1.12) 1.73 (1.18 - 2.52) 0.94 (0.79 - 1.13) 1.43 (0.76 - 2.70) 0.94 (0.90 - 0.98) | P value 0·661 0·110 0·005 0·523 0·226 |
| Indoor Air Pollutant Exposure (N = 429) Site Mbekweni Indoor air pollutant exposure PM10 above ambient standard Infant characteristics Male WAZ at birth Maternal HIV exposure Age in months SES Quartiles Lowest SES | IRR (95% CI) 0.91 (0.61 - 1.37) 0.61 (0.33 - 1.12) 1.73 (1.18 - 2.52) 0.94 (0.79 - 1.13) 1.43 (0.76 - 2.70) 0.94 (0.90 - 0.98) 1.24 (0.73 - 2.11) | P value 0.661 0.110 0.005 0.523 0.226 0.002 |
| Indoor Air Pollutant Exposure (N = 429) Site Mbekweni Indoor air pollutant exposure PM10 above ambient standard Infant characteristics Male WAZ at birth Maternal HIV exposure Age in months SES Quartiles | IRR (95% CI) 0.91 (0.61 - 1.37) 0.61 (0.33 - 1.12) 1.73 (1.18 - 2.52) 0.94 (0.79 - 1.13) 1.43 (0.76 - 2.70) 0.94 (0.90 - 0.98) | P value 0.661 0.110 0.005 0.523 0.226 0.002 |

| Duration of exclusive breast feeding (months) | 0.93 (0.85 - 1.02) | 0.108 |
|---|--------------------|-------|
| Duration of exclusive breast feeding (months) | 0.93 (0.83 - 1.02) | 0.108 |

WAZ, weight-for-age z-score; HIV, human immunodeficiency virus; SES, socio-economic status; PM10, particulate matter

Supplemental Table 7: Multivariable analysis for wheezing and postnatal environmental exposures

| Smoke Exposure | | |
|--|--------------------|---------|
| (N = 875) | IDD (050) CD | D 1 |
| 7.10 | IRR (95% CI) | P value |
| Self-reported smoke exposure | | |
| Maternal smoking | 1.27 (1.03 - 1.56) | 0.024 |
| Any household member smoking | 1.55 (1.17 - 2.06) | 0.002 |
| nfant characteristics | | |
| Male | 1.44 (1.18 - 1.74) | < 0.001 |
| WAZ at birth | 0.96 (0.88 - 1.04) | 0.334 |
| Maternal HIV exposure | 0.58 (0.40 - 0.85) | 0.006 |
| ES quartiles | | |
| Lowest SES | 0.99 (0.73 - 1.35) | 0.973 |
| Low-mod SES | 1.28 (0.97 - 1.70) | 0.079 |
| Mod-high SES | 1.52 (1.16 - 1.99) | 0.002 |
| nfant feeding | | |
| Duration exclusively breast fed in months | 0.98 (0.93 - 1.03) | 0.410 |
| (N = 336) | IDD (050/ CD) | D 1 |
| | IRR (95% CI) | P value |
| ndoor air pollution | | |
| Toluene above ambient standard | 0.60 (0.35 - 1.05) | 0.071 |
| Particulate Matter (PM10) above ambient standard | 0.84 (0.56 - 1.26) | 0.402 |
| Benzene above ambient standard | 1.17 (0.87 - 1.57) | 0.291 |
| nfant characteristics | | |
| Male | 1.45 (1.09 - 1.93) | 0.011 |
| WAZ at birth | 0.91 (0.80 - 1.04) | 0.183 |
| Maternal HIV exposure | 0.98 (0.58 - 1.65) | 0.934 |
| ES quartiles | | |
| Lowest SES | 0.90 (0.57 - 1.42) | 0.646 |
| Low-mod SES | 1.07 (0.70 - 1.64) | 0.758 |
| Mod-high SES | 1.78 (1.20 - 2.64) | 0.004 |
| nfant feeding | | |
| Duration exclusively breast fed in months | 0.89 (0.83 - 0.96) | 0.003 |
| | | |

WAZ, weight-for-age z-score; HIV, human immunodeficiency virus; SES, socio-economic status

Supplemental Table 8: Multivariate analysis for combined antenatal and postnatal environmental exposures

| Wheeze | | |
|---|---|---------|
| | IRR (95% CI) | P value |
| Combined | , | |
| Smoke exposure | 1.79 (1.34 - 2.38) | < 0.001 |
| Infant characteristics | <u> </u> | |
| Male | 1.45 (1.20 - 1.77) | < 0.001 |
| WAZ at birth | 0.95 (0.87 - 1.04) | 0.239 |
| Maternal HIV exposure | 0.50 (0.34 - 0.74) | 0.001 |
| SES quartiles | | |
| Lowest SES | 0.94 (0.70 - 1.28) | 0.698 |
| Low-mod SES | 1.26 (0.95 - 1.67) | 0.106 |
| Mod-high SES | 1.50 (1.15 - 1.97) | 0.003 |
| Infant feeding | | |
| Duration exclusively breast fed in months | 0.98 (0.93 - 1.03) | 0.339 |
| · | | |
| Lower respiratory tract infection (LRTI) | | |
| | Odds Ratio (95% CI) | P value |
| Site | | |
| Mbekweni | 1.19 (0.90 - 1.58) | 0.222 |
| Combined | | |
| Smoke exposure | 1.39 (0.98 - 1.96) | 0.067 |
| Infant characteristics | | |
| Male | 1.74 (1.34 - 2.26) | < 0.001 |
| WAZ at birth | 0.94 (0.84 - 1.06) | 0.304 |
| Maternal HIV exposure | 1.22 (0.83 - 1.80) | 0.310 |
| Age of EPI in months | 0.90 (0.88 - 0.93) | < 0.001 |
| SES quartiles | | |
| Lowest SES | 1.20 (0.82 - 1.75) | 0.348 |
| Low-mod SES | 1.58 (1.10 - 2.26) | 0.013 |
| Mod-high SES | 0.95 (0.65 - 1.39) | 0.796 |
| Infant feeding | | |
| Duration exclusively breast fed in months | 0.96 (0.91 - 1.03) | 0.341 |

WAZ, weight-for-age z-score; HIV, human immunodeficiency virus; EPI, extended programme for immunisation; SES, socio-economic status

Supplemental Table 9: Multivariate analysis for combined environmental tobacco smoke (ETS) and indoor air pollution (IAP) exposures

| Wheeze | | |
|---|---------------------|---------|
| | IRR (95% CI) | P value |
| Combined | | |
| IAP / ETS exposure | 1.96 (1.32 - 2.92) | 0.001 |
| Infant characteristics | | |
| Male | 1.42 (1.17 - 1.73) | <0.001 |
| WAZ at birth | 0.95 (0.87 - 1.03) | 0.234 |
| Maternal HIV exposure | 0.51 (0.34 - 0.75) | 0.001 |
| SES quartiles | | |
| Lowest SES | 0.99 (0.73 - 1.34) | 0.955 |
| Low-mod SES | 1.26 (0.96 - 1.67) | 0.100 |
| Mod-high SES | 1.53 (1.17 - 2.00) | 0.002 |
| Infant feeding | | |
| Duration exclusively breast fed in months | 1.00 (0.93 - 1.03) | 0.365 |
| · | | |
| Lower respiratory tract infection (LRTI) | | |
| | Odds Ratio (95% CI) | P value |
| Site | | |
| Mbekweni | 1.11 (0.84 - 1.46) | 0.454 |
| Combined | | |
| IAP / ETS exposure | 0.99 (0.66 - 1.50) | 0.994 |
| Infant characteristics | | |
| Male | 1.67 (1.30 - 2.16) | 0.000 |
| WAZ at birth | 0.90 (0.83 - 1.05) | 0.250 |
| Maternal HIV exposure | 1.27 (0.87 - 1.87) | 0.218 |
| Age of EPI in months | 0.90 (0.88 - 0.93) | 0.000 |
| SES quartiles | | |
| Lowest SES | 1.29 (0.89 - 1.88) | 0.185 |
| Low-mod SES | 1.64 (1.15 - 2.34) | 0.006 |
| Mod-high SES | 0.99 (0.68 - 1.45) | 0.976 |
| Infant feeding | | |
| Duration exclusively breast fed in months | 0.97 (0.91 - 1.03) | 0.311 |

WAZ, weight-for-age z-score; HIV, human immunodeficiency virus; EPI, extended programme for immunisation; SES, socio-economic status

References

- 1. Stein DJ, Seedat S, Herman AA, Heeringa SG, Moomal H, Myer L, Suliman S, Koza L, Williams D. Findings from the first South African Stress and Health Study 2007.
- 2. Alkire S, Foster J. Understandings and misunderstandings of multidimensional poverty measurement. J Econ Inequal 2011; 9: 289-314.
- 3. Alkire S, Foster J. Counting and multidimensional poverty measurement. J Public Econ 2011; 95: 476-487.
- 4. Vanker A, Barnett W, Nduru PM, Gie RP, Sly PD, Zar HJ. Home environment and indoor air pollution exposure in an African birth cohort study. *Science of the Total Environment* 2015; 536: 362-367.
- 5. Government Gazette Republic of South Africa. National Ambient Air Quality Standards. 2009 [cited 2014 August]. Available from: https://www.environment.gov.za/sites/default/files/legislations/nemaqa_airquality_g32816gon1210.pdf.
- 6. Vanker A, Barnett W, Brittain K, Gie RP, Koen N, Myers B, Stein DJ, Zar HJ. Antenatal and early life tobacco smoke exposure in an African birth cohort study. *Int J Tuberc Lung Dis* 2016; 20: 729-737.
- 7. Siemens. Immulite 1000 Nicotine Metabolite [pamphlet]. 2009 25-08-2009 [cited 2014 November]. Available from: http://www.healthcare.siemens.com/immunoassay/systems/immulite-1000-im.
- 8. World Health Organisation. Integrated management of childhood illness: distance learning course. . Geneva; 2014.
- 9. Zar HJ, Barnett W, Stadler A, Gardner-Lubbe S, Myer L, Nicol MP. Aetiology of childhood pneumonia in a well vaccinated South African birth cohort: a nested case-control study of the Drakenstein Child Health Study. *The Lancet Respiratory medicine* 2016; 4: 463-472.
- 10. le Roux DM, Myer L, Nicol MP, Zar HJ. Incidence of childhood pneumonia: facility-based surveillance estimate compared to measured incidence in a South African birth cohort study. *BMJ Open* 2015; 5: e009111.
- 11. Zar HJ, Barnett W, Myer L, Stein DJ, Nicol MP. Investigating the early-life determinants of illness in Africa: the Drakenstein Child Health Study. *Thorax* 2015; 70: 592-594.
- 12. Fenton TR, Nasser R, Eliasziw M, Kim JH, Bilan D, Sauve R. Validating the weight gain of preterm infants between the reference growth curve of the fetus and the term infant. *BMC pediatrics* 2013; 13: 92.